## Amendments to the Claims

Please cancel Claims 7-10. Please amend Claims 2-6, 12, 14-20, 22-24, 26, 27, 29-34, 36-38 and 45. Please add new Claims 46-49. The Claim Listing below will replace all prior versions of the claims in the application:

## **Claim Listing**

- 1. (Original) An antibody or antigen-binding fragment thereof which binds to a mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor, wherein said antibody or antigen-binding fragment thereof inhibits binding of a ligand to the receptor.
- 2. (Currently amended) The An antibody or antigen-binding fragment according to Claim 1, wherein said antibody or antigen-binding fragment thereof inhibits one or more functions associated with binding of the ligand to the receptor.
- 3. (Currently amended) The An antibody or antigen-binding fragment thereof according to Claim 1, wherein the mammalian CC-chemokine receptor 4 (CCR4) is a human CC-chemokine receptor 4 (CCR4).
- 4. (Currently amended) <u>The An</u> antibody or antigen-binding fragment thereof according to Claim 1, wherein the antibody is selected from the group consisting of:
  - a) monoclonal antibody 1G1;
  - b) an antibody having an epitopic specificity which is the same as or similar to that of 1G1;
  - b) c) an antibody which can compete with 1G1 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - c) d) monoclonal antibody 2B10;
  - e) an antibody having an epitopic specificity which is the same as or similar to that of 2B10:
  - <u>d)</u> f) an antibody which can compete with 2B10 for binding to mammalian CC-chemokine receptor 4 (CCR4);

- e) g) monoclonal antibody 10E4;
- h)——an antibody having an epitopic specificity which is the same as or similar to that of 10E4;
- <u>f</u>) i) an antibody which can compete with 10E4 for binding to mammalian CC-chemokine receptor 4 (CCR4); and
- g) j) an antigen-binding fragments fragment of any one of (a) through (i) (f) which bind binds to mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof.
- 5. (Currently amended) <u>The An</u> antibody or antigen-binding fragment thereof according to Claim 1, wherein the ligand is a chemokine.
- 6. (Currently amended) <u>The An</u> antibody or antigen-binding fragment thereof according to Claim 5, wherein the chemokine is any one or more of TARC, MDC, MCP-1, MIP-1α and RANTES.

## 7-10. (Canceled)

- 11. (Original) A test kit for use in detecting the presence of a mammalian CC-chemokine receptor 4 (CCR4) or portion thereof in a biological sample comprising
  - a) at least one antibody or antigen-binding fragment thereof which binds to a mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor, wherein said antibody or antigen-binding fragment thereof inhibits binding of a ligand to the receptor; and
  - b) one or more ancillary reagents suitable for detecting the presence of a complex between said antibody or antigen-binding fragment thereof and said mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof.
- 12. (Currently amended) The A test kit according to Claim 11, wherein the antibody is selected from the group consisting of:
  - i) monoclonal antibody 1G1;

- ii) an antibody having an epitopic specificity which is the same as or similar to that of 1G1;
- ii) iii) an antibody which can compete with 1G1 for binding to mammalian CC-chemokine receptor 4 (CCR4);
- iii) iv) monoclonal antibody 2B10;
- v) an antibody having an epitopic specificity which is the same as or similar to that of 2B10;
- iv) vi) an antibody which can compete with 2B10 for binding to mammalian CC-chemokine receptor 4 (CCR4);
- v) vii) monoclonal antibody 10E4;
- viii) an antibody having an epitopic specificity which is the same as or similar to that of 10E4;
- vi) ix) an antibody which can compete with 10E4 for binding to mammalian CC-chemokine receptor 4 (CCR4); and
- vii) x) an antigen-binding fragments fragment of any one of (i) through (ix) (vi) which bind binds to mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof [[.]]; and
- viii) combinations of the foregoing.
- 13. (Original) A method of inhibiting the interaction of a cell bearing mammalian CC-chemokine receptor 4 (CCR4) with a ligand thereof, comprising contacting said cell with an effective amount of an antibody or antigen-binding fragment thereof which binds to mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor and inhibits binding of said ligand to the receptor.
- 14. (Currently amended) <u>The A</u> method according to Claim 13, wherein the cell is selected from the group consisting of lymphocytes, monocytes, granulocytes, T cells, basophils, and cells comprising a recombinant nucleic acid encoding CCR4 or a portion thereof.

- 15. (Currently amended) The A method according to Claim 14, wherein the T cells are selected from the group consisting of CD8+ cells, CD25+ cells, CD4+ cells and CD45RO+ cells.
- 16. (Currently amended) <u>The A method according to Claim 13</u>, wherein the ligand is a chemokine.
- 17. (Currently amended) <u>The A method according to Claim 16</u>, wherein the chemokine is any one or more of TARC, MDC, MCP-1, MIP-1α and RANTES.
- 18. (Currently amended) The A method according to Claim 13, wherein the antibody or antigen-binding fragment thereof is selected from the group consisting of:
  - a) monoclonal antibody 1G1;
  - b) an antibody having an epitopic specificity which is the same as or similar to that of 1G1;
  - <u>b)</u> c) an antibody which can compete with 1G1 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - c) d) monoclonal antibody 2B10;
  - e) an antibody having an epitopic specificity which is the same as or similar to that of 2B10;
  - <u>d)</u> f) an antibody which can compete with 2B10 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - e) g) monoclonal antibody 10E4;
  - h) an antibody having an epitopic specificity which is the same as or similar to that of 10E4;
  - <u>f) i)</u> an antibody which can compete with 10E4 for binding to mammalian CC-chemokine receptor 4 (CCR4); and
  - <u>an</u> antigen-binding fragments fragment of any one of (a) through (i) (f) which bind binds to mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof[[.]]:

    and

## h) combinations of the foregoing.

- 19. (Currently amended) A method of detecting expression of mammalian CC-chemokine receptor 4 (CCR4) or portion thereof by a cell or fraction of said cell, comprising:
  - a) contacting a composition comprising a cell or fraction of said cell to be tested with an antibody or antigen-binding fragment thereof which binds to mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor and inhibits binding of said a ligand to the receptor, under conditions appropriate for binding of said antibody or antigen-binding fragment thereof to a mammalian CCR4 or portion thereof; and
  - b) detecting binding of said antibody or antigen-binding fragment thereof, wherein the binding of said antibody or antigen-binding fragment thereof indicates the presence of said receptor or portion of said receptor on said cell.
- 20. (Currently amended) The A method according to Claim 19, wherein the antibody or antigen-binding fragment thereof is selected from the group consisting of:
  - i) monoclonal antibody 1G1;
  - ii) an antibody having an epitopic specificity which is the same as or similar to that of 1G1;
  - ii) iii) an antibody which can compete with 1G1 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - iii) iv) monoclonal antibody 2B10;
  - v) an antibody having an epitopic specificity which is the same as or similar to that of 2B10;
  - iv) vi) an antibody which can compete with 2B10 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - vii) antigen-binding fragments of any one of (i) through (vi) which bind mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof;
  - v) viii) monoclonal antibody 10E4;

- ix) an antibody having an epitopic specificity which is the same as or similar to that of 10E4;
- vi) x) an antibody which can compete with 10E4 for binding to mammalian CC-chemokine receptor 4 (CCR4); and
- vii) an antigen binding fragment of any one of (i) through (vi) which binds
  mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof; and
  viii xi) combinations of the foregoing.
- 21. (Original) The method according to Claim 20, wherein the composition is a sample comprising human cells.
- 22. (Currently amended) A method of detecting a mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor, comprising:
  - a) contacting a sample to be tested with an antibody or antigen-binding fragment thereof which binds to mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor and inhibits binding of said a ligand to the receptor under conditions appropriate for binding of said antibody or fragment thereof to said mammalian CCR4 or portion thereof; and
  - b) detecting or measuring binding of said antibody or antigen-binding fragment thereof,

wherein the binding of said antibody or antigen-binding fragment thereof to material in said sample is indicative of the presence of a mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor in said sample.

- 23. (Currently amended) The A method according to Claim 22, wherein the antibody or antigeni antigen-binding fragment thereof is is selected from the group consisting of:
  - i) monoclonal antibody 1G1;
  - ii) an antibody having an epitopic specificity which is the same as or similar to that of 1G1;

- ii) iii) an antibody which can compete with 1G1 for binding to mammalian CC-chemokine receptor 4 (CCR4);
- iii) iv) monoclonal antibody 2B10;
- v) an antibody having an epitopic specificity which is the same as or similar to that of 2B10;
- vi) vi) an antibody which can compete with 2B10 for binding to mammalian CC-chemokine receptor 4 (CCR4);
- v) vii) monoclonal antibody 10E4;
- viii) an antibody having an epitopic specificity which is the same as or similar to that of 10E4;
- vi) ix) an antibody which can compete with 10E4 for binding to mammalian CC-chemokine receptor 4 (CCR4);
- vii x) an antigen-binding fragments fragment of any one of (i) through (ix) (vi) which bind binds to mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof; and
- viii xi) combinations of the foregoing.
- 24. (Currently amended) The A method according to Claim 22, wherein the sample is a cellular fraction which, in normal individuals, comprises a mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor.
- 25. (Original) A method of inhibiting a function associated with binding of a chemokine to a mammalian CC-chemokine receptor 4 (CCR4) or a functional portion of said receptor, comprising contacting a composition comprising the receptor or portion with an effective amount of an antibody or antigen-binding fragment thereof which binds to a mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor, wherein said antibody or fragment inhibits binding of said chemokine to mammalian CC-chemokine receptor 4 (CCR4) and inhibits one or more functions associated with binding of the chemokine to the receptor.

- 26. (Currently amended) <u>The A method according to Claim 25</u>, wherein the chemokine is any one or more of TARC, MDC, MCP-1, MIP-1α and RANTES.
- 27. (Currently amended) The A method according to Claim 25, wherein the antibody or antigen-binding fragment is selected from the group consisting of:
  - a) monoclonal antibody 1G1;
  - b) an antibody having an epitopic specificity which is the same as or similar to that of 1G1;
  - b) c) an antibody which can compete with 1G1 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - c) d) monoclonal antibody 2B10;
  - e) an antibody having an epitopic specificity which is the same as or similar to that of 2B10;
  - <u>d)</u> f) an antibody which can compete with 2B10 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - e) g) monoclonal antibody 10E4;
  - h) an antibody having an epitopic specificity which is the same as or similar to that of 10E4;
  - <u>f)</u> i) an antibody which can compete with 10E4 for binding to mammalian CC-chemokine receptor 4 (CCR4); and
  - g) j) an antigen-binding fragments fragment of any of (a) through (i) (f) which bind binds to mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof[[.]]; and
  - h) combinations of the foregoing.
- 28. (Original) A method of detecting or identifying an agent which binds a mammalian CC-chemokine receptor 4 (CCR4) or ligand-binding variant thereof, comprising combining
  - a) an agent to be tested;
  - b) an antibody or antigen-binding fragment which binds to a mammalian CCchemokine receptor 4 (CCR4) or portion of said receptor, wherein said antibody

- or antigen-binding fragment thereof inhibits binding of a ligand to the receptor; and
- c) a composition comprising a mammalian CC-chemokine receptor 4 (CCR4) or a ligand-binding variant thereof,

under conditions suitable for binding of said antibody or antigen-binding fragment to said mammalian CC-chemokine receptor 4 (CCR4) or ligand-binding variant thereof, and detecting or measuring binding of said antibody or antigen-binding fragment to said mammalian CC-chemokine receptor 4 (CCR4) or ligand-binding variant thereof.

- 29. (Currently amended) The A method according to Claim 28, wherein the antibody or antigen-binding fragment thereof is selected from the group consisting of:
  - a) monoclonal antibody 1G1;
  - an antibody having an epitopic specificity which is the same as or similar to that of 1G1;
  - b) c) an antibody which can compete with 1G1 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - c) d) monoclonal antibody 2B10;
  - e) an antibody having an epitopic specificity which is the same as or similar to that of 2B10;
  - d) f) an antibody which can compete with 2B10 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - e) g) monoclonal antibody 10E4;
  - h) an antibody having an epitopic specificity which is the same as or similar to that of 10E4;
  - <u>f)</u> i) an antibody which can compete with 10E4 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - g) j) an antigen-binding fragments fragment of any of (a) through (i) (f) which bind binds to mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof; and
  - $\underline{h}$  combinations of the foregoing.

- 30. (Currently amended) The A method according to Claim 28, wherein the formation of a complex between said antibody or antigen-binding fragment and said mammalian CC-chemokine receptor 4 (CCR4) or ligand-binding variant is monitored, and wherein a decrease in the amount of complex formed relative to a suitable control is indicative that the agent binds said receptor or ligand-binding variant thereof.
- 31. (Currently amended) The A method according to Claim 28, wherein the composition comprising a mammalian CC-chemokine receptor 4 (CCR4) or a ligand-binding variant thereof is a cell bearing recombinant CC-chemokine receptor 4 (CCR4) or ligand-binding variant thereof.
- 32. (Currently amended) The A method according to Claim 28, wherein the composition comprising a mammalian CC-chemokine receptor 4 (CCR4) or a ligand-binding variant thereof is a membrane fraction of said cell bearing recombinant CC-chemokine receptor 4 (CCR4) or ligand-binding variant thereof.
- 33. (Currently amended) The A method according to Claim 28, wherein the antibody or antigen-binding fragment thereof is labeled with a label selected from the group consisting of a radioisotope, spin label, antigen label, enzyme label, fluorescent group and chemiluminescent group.
- 34. (Currently amended) The A method according to Claim 28, wherein the agent is an antibody having specificity for a mammalian CC-chemokine receptor 4 (CCR4) or antigen-binding fragment thereof.
- 35. (Original) A method of inhibiting leukocyte trafficking in a patient, comprising administering to the patient a composition comprising an effective amount of an antibody or antigen-binding fragment thereof which binds to a mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor and inhibits binding of a ligand to the receptor.

- 36. (Currently amended) The A method according to Claim 35, wherein the ligand is a chemokine.
- 37. (Currently amended) <u>The A method according to Claim 36</u>, wherein the chemokine is any one or more of TARC, MDC, MCP-1, MIP-1α and RANTES.
- 38. (Currently amended) The A method according to Claim 35, wherein the antibody or antigen-binding fragment thereof is selected from the group consisting of:
  - a) monoclonal antibody 1G1;
  - b) an antibody having an epitopic specificity which is the same as or similar to that of 1G1;
  - b) c) an antibody which can compete with 1G1 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - c) d) monoclonal antibody 2B10;
  - e) an antibody having an epitopic specificity which is the same as or similar to that of 2B10;
  - <u>d)</u> f) an antibody which can compete with 2B10 for binding to mammalian CC-chemokine receptor 4 (CCR4);
  - e) g) monoclonal antibody 10E4;
  - h) an antibody having an epitopic specificity which is the same as or similar to that of 10E4;
  - <u>f)</u> i) an antibody which can compete with 10E4 for binding to mammalian CC-chemokine receptor 4 (CCR4); and
  - g) j) an antigen-binding fragments fragment of any of (a) through (f) which bind binds to mammalian CC-chemokine receptor 4 (CCR4) or a portion thereof[[.]]; and
  - <u>h)</u> combinations of the foregoing.
- 39. (Original) A composition comprising an antibody or antigen-binding fragment thereof which binds to a mammalian CC-chemokine receptor 4 (CCR4) or portion of said

- receptor, wherein said antibody or antigen-binding fragment thereof inhibits binding of a ligand to the receptor, and an optional physiologically acceptable vehicle.
- 40. (Original) An antibody or antigen-binding fragment thereof which binds to a mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor, wherein said antibody or antigen-binding fragment thereof inhibits binding of a ligand to the receptor with an IC<sub>50</sub> of less than about 1.5 μg/ml.
- 41. (Original) An antibody or antigen-binding fragment thereof which binds to a mammalian CC-chemokine receptor 4 (CCR4) or portion of said receptor, wherein said antibody or antigen-binding fragment thereof inhibits binding of a ligand to the receptor with an IC<sub>50</sub> of less than about 1.5 ng/ml.
- 42. (Original) A method of treating a CC-chemokine receptor 4 (CCR4)-mediated disorder in a patient, comprising administering to the patient an effective amount of an antibody or antigen-binding fragment thereof which binds to mammalian CC-chemokine receptor 4 (CCR4) or portion thereof.
- 43. (Original) A method of treating an inflammatory disorder in a patient, comprising administering to the patient an effective amount of an antibody or antigen-binding fragment thereof which binds to mammalian CC-chemokine receptor 4 (CCR4) or portion thereof.
- 44. (Original) The 10E4 hybridoma cell line deposited under ATCC Accession No. PTA-1203.
- 45. (Currently amended) A monoclonal antibody produced by the <u>10E4</u> hybridoma cell line according to Claim 44 deposited under ATCC Accession No. PTA-1203 or an antigenbinding fragment thereof.

- 46. (New) The antibody or antigen-binding fragment thereof according to Claim 6, wherein the chemokine is TARC or MDC.
- 47. (New) The method according to Claim 17, wherein the chemokine is TARC or MDC.
- 48. (New) The method according to Claim 26, wherein the chemokine is TARC or MDC.
- 49. (New) The method according to Claim 37, wherein the chemokine is TARC or MDC.